TCTAGACTGGACAGCATCCACAAGAGAAGCACCTAGAAGGAGAATTTTCCCCAGCAGCTTGCTCAGGACCC
TGCAGGAGCCGCAGCTGGGACTGGACCTGCTGTTAACCATGAACTCTTCCTGCTGCCTGTCTTCTGTTTCT
CCGATGCTGCCTAACCTCTCTGAGCACCCTGCAGCCCCTCCTGCCAGCAACCGGAGCGGCAGTGGGTTCTG
TGAGCAGGTCTTCATCAAGCCGGAGGTCTTCCTGGCTCTGGGCATCGTCAGTCTGATGGAAAACATCCTGG
TGATCCTGGCTGTGGTCAGGAATGGCAACCTGCACTCTCCCATGTACTTCTTCCTGTGCAGCCTGGCTGCA
GCCGACATGCTGGTGAGCCTGTCCAACTCCCTGGAGACCATCATGATCGCCGTGATCAACAGCGACTCCCT
GACCTTGGAGGACCAGTTTATCCAGCACATGGATAATATCTTCGACTCTATGATTTGCATCTCCCTGGTGG
CCTCCATCTGCAACCTCCTGGCCATTGCCATCGACAGGTACGTCACCATCTTCTATGCCCTTCGGTACCAC
AGCATCATGACAGTTAGGAAAGCCCTCACCTTGATCGGGGTCATCTGGGTCTGCTGCGGCATCTGCGGCGT
GATGTTCATCATCTACTCCGAGAGCAAGATGGTCATCGTGTGTCTCATCACCATGTTCTTCGCCATGGTGC
TCCTCATGGGCACCCTATATATCCACATGTTCCTCTTCGCCAGGCTCCACGTCCAGCGCATCGCAGTGCTG
CCCCCTGCTGGCGTGGTGGCCCCACAGCAGCACTCCTGCATGAAGGGGGCTGTCACCATCACTATCCTGCT
GGGTGTTTTCATCTTCTGCTGGGCGCCTTTCTTCCTCCACCTGGTCCTCATCATCACCTGCCCCACCAATC
CCTACTGCATCTGCTACACGGCCCATTTCAACACCTACCTGGTTCTCATCATGTGCAACTCCGTCATCGAC
CCCCTCATCTACGCCTTCCGCAGCCTGGAGCTGCGCAACACGTTCAAGGAGATTCTCTGCGGCTGCAACAG
CATGAACTTGGGCTAGGATGCCCGTGGAGGTGTTCCACATCCAGCCAAGAGACAAAAACAACGCTCAGACG
GGACGTAAAAGGGTGTTAGGAGCTGGAACTGTGCTTGGCTTCGTCTGTAAGCTCGTGGCCCTTTGCAGACG
GGACACGGCGTAGGATGGGCTGTCTGTGAGGATCTGTGTGTGGGTAAGTCAGTTTGATCTAGCACATAGCC
TGGAAGAATCAGGCAAAGCAGCCCTGAGTGTCATCTGTGTTCATTGCTAGGCACCCAGGGTTTGTGGCCCC
TGCCTGCTTATTGGCTTTGTACCAGTAACTGTGCTTCAAGCCAACCAGACCGGAGGGCTCTCGTGAGCAGA
AAGAGTGCTTAGACTTCCGGCAAGCATCCTGGCTCACAGCGGCCACCTCCTGACCACTACCGGGAGAGCTT
TGCACATATTCTGTGGGAGATTGAGTGAAGCCCTGAAAACAATGTGATATTTGCTGCTCCCTTCCAGAACT
TACATCTGTGCCAGCCTCCCCGAACCCCTGCACAGAGACATGACCCCCTTCTCCCTGTGCCGTTGTCATGG
TTGTTATTATTGTTGGAGTTTTGTTCGTTAAAATCTAAGCTT (SEQ ID NO:1)

MNSSCCLSSVSPMLPNLSEHPAAPPASNRSGSGFCEQVFIKPEVFLALGIVSLMENILVILAVVRNGNLHS
PMYFFLCSLAAADMLVSLSNSLETIMIAVINSDSLTLEDQFIQHMDNIFDSMICISLVASICNLLAIAIDR
YVTIFYALRYHSIMTVRKALTLIGVIWVCCGICGVMFIIYSESKMVIVCLITMFFAMVLLMGTLYIHMFLF
ARLHVQRIAVLPPAGVVAPQQHSCMKGAVTITILLGVFIFCWAPFFLHLVLIITCPTNPYCICYTAHFNTY
LVLIMCNSVIDPLIYAFRSLELRNTFKEILCGCNSMNLG (SEQ ID NO:2)

FIGURE 1

Underlined = deleted in targeting construct

Bold = sequence flanking Neo insert in targeting construct

TCTAGACTGGACAGCATCCACAAGAGAAGCACCTAGAAGGAGAATTTTCCCCAGCAGCTT
GCTCAGGACCCTGCAGGAGCCGCAGCTGGGACTGGACCTGCTGTTAACCATGAACTCTTC
CTGCTGCCTGTCTTCTGTTTCTCCGATGCTGCCTAACCTCTCTGAGCACCCTGCAGCCCC
TCCTGCCAGCAACCGGAGCGGCAGTGGGTTCTGTGAGCAGGTCTTCATCAAGCCGGAGGT
CTTCCTGGCTCTGGGCATCGTCAGTCTGATGGAAAACATCCTGGTGATCCTGGCT<u>GTGGT</u>
<u>CAGGAATGGCAACCTGCACTCTCCCATGTACTTCTTCCTGTGCAGCCTGGCTGCAGCCGA</u>
<u>CATGCTGGTGAGCCTGTCCAACTCCCTGGAGACCATCATGATCGCCGTGATCAACAGCGA</u>
<u>CTCCCTGACCTTGGAGGACCAGTTTATCCAGCACATGGATAATATCTTCGACTCTATGAT</u>
<u>TTGCATCTCCCTGGTGGCCTCCATCTGCAACCTCCTGGC</u>**CATTGCCATCGACAGGTACGT**
CACCATCTTCTATGCCCTTCGGTACCACAGCATCATGACAGTTAGGAAAGCCCTCACCTT
GATCGGGGTCATCTGGGTCTGCTGCGGCATCTGCGGCGTGATGTTCATCATCTACTCCGA
GAGCAAGATGGTCATCGTGTGTGTCTCATCACCATGTTCTTCGCCATGGTGCTCCTCATGGG
CACCCTATATATCCACATGTTCCTCTTCGCCAGGCTCCACGTCCAGCGCATCGCAGTGCT
GCCCCCTGCTGGCGTGGTGGCCCCACAGCAGCACTCCTGCATGAAGGGGGCTGTCACCAT
CACTATCCTGCTGGGTGTTTTCATCTTCTGCTGGGCGCCTTTCTTCCTCCACCTGGTCCT
CATCATCACCTGCCCCACCAATCCCTACTGCATCTGCTACACGGCCCATTTCAACACCTA
CCTGGTTCTCATCATGTGCAACTCCGTCATCGACCCCCTCATCTACGCCTTCCGCAGCCT
GGAGCTGCGCAACACGTTCAAGGAGATTCTCTGCGGCTGCAACAGCATGAACTTGGGCTA
GGATGCCCGTGGAGGTGTTCCACATCCAGCCAAGAGACAAAAACAACGCTCAGACGGGAC
GTAAAAGGGTGTTAGGAGCTGGAACTGTGCTTGGCTTCGTCTGTAAGCTCGTGGCCCTTT
GCAGACGGGACACGGCGTAGGATGGGCTGTCTGTGAGGATCTGTGTGTGGGTAAGTCAGT
TTGATCTAGCACATAGCCTGGAAGAATCAGGCAAAGCAGCCCTGAGTGTCATCTGTGTTC
ATTGCTAGGCACCCAGGGTTTGTGGCCCCTGCCTGCTTATTGGCTTTGTACCAGTAACTG
TGCTTCAAGCCAACCAGACCGGAGGGCTCTCGTGAGCAGAAAGAGTGCTTAGACTTCCGG
CAAGCATCCTGGCTCACAGCGGCCACCTCCTGACCACTACCGGGAGAGCTTTGCACATAT
TCTGTGGGAGATTGAGTGAAGCCCTGAAAACAATGTGATATTTGCTGCTCCCTTCCAGAA
CTTACATCTGTGCCAGCCTCCCCGAACCCCTGCACAGAGACATGACCCCCTTCTCCCTGT
GCCGTTGTCATGGTTGTTATTATTGTTGGAGTTTTGTTCGTTAAAATCTAAGCTT
(SEQ ID NO:1)

FIGURE 2A

Gene Sequence
Structure *

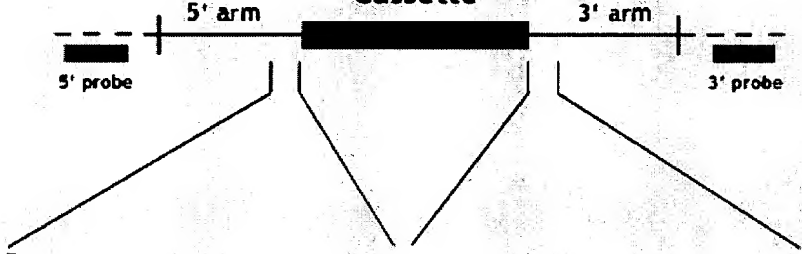296 bp   Sequence Deleted   519 bp

Size of CDS: 1675 bp

Targeting Vector* (genomic sequence)

Construct Number: 2787

LacZ-Neo
Cassette

5' arm   3' arm

5' probe   3' probe

Arm Length:
5': 1.6 kb
3': 4 kb

────── Targeting Vector
- - - - - Endogenous Locus

* Not drawn to scale

5'>ACCTGCTGTTAACCATGAACT
CTTCCTGCTGCCTGTCTTCTGTTT
CTCCGATGCTGCCTAACCTCTCTG
AGCACCCTGCAGCCCCTCCTGCCA
GCAACCGGAGCGGCAGTGGGTTCT
GTGAGCAGGTCTTCATCAAGCCGG
AGGTCTTCCTGGCTCTGGGCATCG
TCAGTCTGATGGAAAACATCCTGG
TGATCCTGGCT<3'
(SEQ ID NO:3)

5'>CATTGCCATCGACAGGTACGT
CACCATCTTCTATGCCCTTCGGTA
CCACAGCATCATGACAGTGAGGAA
AGCCCTCACCTTGATCGGGGTCAT
CTGGGTCTGCTGCGGCATCTGCGG
CGTGATGTTCATCATCTACTCCGA
GAGCAAGATGGTCATCGTGTGTCT
CATCACCATGTTCTTCGCCATGGT
GCTCCTCATGG<3'
(SEQ ID NO:4)

FIGURE 2B